DOCKET NO.: ISIS-4682

Application No.: 09/775,967

Office Action Dated: June 11, 2003

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

A method comprising reacting a nucleoside phosphoramidite with a support bound 1 (original).

oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one

unprotected internucleoside linkage selected from the group consisting of phosphate linkages,

phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an

aromatic heterocyclic amine, a guanidine, or a salt of formula D+E- wherein:

D⁺ is a quaternary tetraalkylammonium cation, or a protonated form of

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an

aromatic heterocyclic amine, or a guanidine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or

unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate

anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate

anion.

The method of claim 1 wherein said neutralizing agent is a salt of formula D⁺ E⁻. 2 (original).

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3 (original). The method of claim 2 wherein E is a tetrazolide anion.

4 (original). The method of claim 1 wherein E is 1H-tetrazolide anion, 5-methylthio-1H-

tetrazolide anion, 5-ethylthio-1H-tetrazolide anion or 1-phenyl-5-thiol-1H-tetrazolide anion.

5 (original). The method of claim 1 wherein E is 1H-tetrazolide anion.

6 (original). The method of claim 3 wherein D⁺ is a protonated form of any of an alkyl, alkenyl

or alkynyl amine having from one to about 20 carbons, an aliphatic heterocyclic amine, an aromatic

heterocyclic amine, or a guanidine.

7-10 (canceled).

11 (currently amended). The method of claim 3 A method comprising reacting a

nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said

support bound oligomer having at least one unprotected internucleoside linkage selected from the group

consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is a salt of formula D+E wherein:

D⁺ is a protonated form of an aromatic heterocyclic amine; and

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E is a tetrazolide anion.

12 (currently amended). The method of claim 3 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is a salt of formula D+E wherein:

D⁺ is a protonated form of a mono-, di- or trialkyl pyridine that is optionally substituted with an amino group; and

E is a tetrazolide anion.

13 (currently amended). The method of claim 3 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is a salt of formula D⁺E⁻ wherein:

D⁺ is a protonated form of any of 2,4,6-collidine, 2,6-lutidine, pyridine, 2-methylpyridine, 2,6-di(t-butyl)pyridine, 4-methyl-2,6-di(t-butyl)pyridine, or 2,4,6-tri(t-butyl)pyridine; and

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E is a tetrazolide anion.

14 (currently amended). The method of claim 3 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is a salt of formula D+E wherein:

D⁺ is a protonated form of an alkylamino substituted pyridine; and E⁻ is a tetrazolide anion.

15 (currently amended). The method of claim 3 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is a salt of formula D+E-wherein:

D⁺ is a protonated form of 4-dimethylaminopyridine; and E⁻ is a tetrazolide anion.

16-20 (canceled).

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21 (original). The method of claim 3 wherein E is 1H-tetrazolide anion.

22-35 (canceled).

36 (currently amended). The method of claim 1 wherein A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, a guanidine, or a salt of formula D+E wherein:

D⁺ is a protonated form of an aromatic heterocyclic amine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

37 (currently amended). The method of claim 1 A method comprising reacting a nucleoside

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phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic amine, an aromatic heterocyclic amine, a guanidine, or a salt of formula D+E- wherein:

D⁺ is a protonated form of a mono-, di- or trialkyl pyridine that is optionally substituted with an amino group; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

The method of claim 1 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an

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aromatic heterocyclic amine, a guanidine, or a salt of formula D+E wherein:

D⁺ is a protonated form of any of 2,4,6-collidine, 2,6-lutidine, pyridine, 2-methylpyridine, 2,6-diethylpyridine, 2,6-di(t-butyl)pyridine, 4-methyl-2,6-di(t-butyl)pyridine, or 2,4,6-tri(t-butyl)pyridine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

The method of claim 1 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, a guanidine, or a salt of formula D⁺E⁻ wherein:

D⁺ is a protonated form of an alkylamino substituted pyridine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate

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anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

40 (currently amended). The method of claim 1 A method comprising reacting a nucleoside phosphoramidite with a support bound oligomer in the presence of a neutralizing agent, said support bound oligomer having at least one unprotected internucleoside linkage selected from the group consisting of phosphate linkages, phosphorothioate linkages, and phosphorodithioate linkages;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, aromatic heterocyclic amine, a guanidine, or a salt of formula D+E wherein:

D⁺ is a protonated form of 4-dimethylaminopyridine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

41-46 (canceled).

47 (original). The method of claim 3 wherein D⁺ is a protonated form of trimethyl amine,

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triethyl amine, triisopropyl amine, tributyl amine, triamyl amine, isopropyldimethyl amine, t-butyldimethyl amine, diisopropylethyl amine, N,N,N',N'-tetramethyl-1,2-diaminoethane, DBU, N-methylmorpholine, N-methylpyrrolidine, N-methylpiperidine, N,N'-dimethylpiperazine, N-ethylpyrrolidine, N-ethylpiperidine, N,N'-diethylpiperazine, 1,5-diazabicyclo[4.3.0]non-5-ene, 1,4-diazabicyclo[2.2.2]octane, or 1,5,7-triazabicyclo[4.4.0]dec-5ene, 2,4,6-collidine, 2,6-lutidine, pyridine, 2-methylpyridine, 2,6-diethylpyridine, 2,6-di(t-butyl)pyridine, 4-methyl-2,6-di(t-butyl)pyridine, or 2,4,6-tri(t-butyl)pyridine, 4-dimethylaminopyridine, or N,N,N'N'-tetramethylguanidine, or tetramethylammonium, tetraethylammonium, tetraethylammonium, tetrabutylammonium, tetrabutylammonium, tetrabutylammonium,

E is 1H-tetrazolide anion, 4,5-dicyanoimidazolide anion, methylsulfonate anion, trifluoromethylsulfonate anion, methylphenylsulfonate anion, trifluoromethylphenylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or trifluoroacetate anion.

48 (currently amended). A method of forming an internucleoside linkage comprising reacting a phosphoramidite of formula:

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$$R_1$$
— O
 R_2
 R_2

wherein:

L₁ is an internucleoside linkage;

 n_1 is 0 to about 100;

R₁ is a hydroxyl protecting group;

R₂ is a 2'-substituent group;

R₄ and R₅ are each independently alkyl having from 1-to about 10 carbon atoms, or R₄ and R₅ taken together with the nitrogen atom to which they are attached form a heterocycle;

B is a nucleobase;

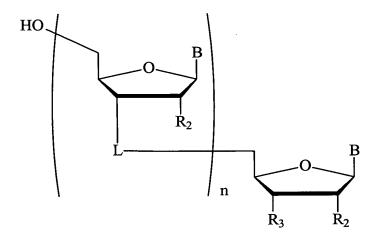
Q is O or S;

Pg is a phosphoryl protecting group;

with a compound of formula:

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wherein

 R_3 is a linker connected to a solid support;

n is from 1 to 100; and

L is an internucleoside linkage of formula:

wherein:

Z is O or S;

X is O or S; and

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Y is a phosphoryl protecting group or a negative charge; provided that at least one Y is a negative charge; wherein said reaction is performed in the presence of a neutralizing agent; wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, a guanidine, or a salt of formula D+E- wherein:

D⁺ is a quaternary tetraalkylammonium cation, or a protonated form of an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, or a guanidine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

49 (original). The method of claim 48 wherein said neutralizing agent is a salt of formula D^+ E^- .

50 (original). The method of claim 49 wherein E is a tetrazolide anion.

51 (original). The method of claim 48 wherein E⁻ is 1H-tetrazolide anion, 5-methylthio-1H-tetrazolide anion, 5-ethylthio-1H-tetrazolide anion or 1-phenyl-5-thiol-1H-tetrazolide anion.

52 (original). The method of claim 48 wherein E is 1H-tetrazolide anion.

53-55 (canceled).

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56 (original). The method of claim 50 wherein D⁺ is a protonated form of an aromatic heterocyclic amine.

57 (original). The method of claim 50 wherein D⁺ is a protonated form of a mono-, di- or trialkyl pyridine that is optionally substituted with an amino group.

58 (original). The method of claim 50 wherein D⁺ is a protonated form of any of 2,4,6-collidine, 2,6-lutidine, pyridine, 2-methylpyridine, 2,6-diethylpyridine, 2,6-di(t-butyl)pyridine, 4-methyl-2,6-di(t-butyl)pyridine, or 2,4,6-tri(t-butyl)pyridine.

59 (original). The method of claim 50 wherein D⁺ is a protonated form of an alkylamino substituted pyridine.

60 (original). The method of claim 50 wherein D^+ is a protonated form of 4-dimethylaminopyridine.

61-65 (canceled).

66 (original). The method of claim 50 wherein E is 1H-tetrazolide anion.

67-80 (canceled).

81 (original). The method of claim 48 wherein D⁺ is a protonated form of an aromatic heterocyclic amine.

82 (original). The method of claim 48 wherein D⁺ is a protonated form of a mono-, di- or

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trialkyl pyridine that is optionally substituted with an amino group.

83 (original). The method of claim 48 wherein D⁺ is a protonated form of any of 2,4,6-collidine, 2,6-lutidine, pyridine, 2-methylpyridine, 2,6-diethylpyridine, 2,6-di(t-butyl)pyridine, 4-methyl-2,6-di(t-butyl)pyridine, or 2,4,6-tri(t-butyl)pyridine.

84 (original). The method of claim 48 wherein D⁺ is a protonated form of an alkylamino substituted pyridine.

85 (original). The method of claim 48 wherein D⁺ is a protonated form of 4-dimethylaminopyridine.

86-91 (canceled).

92 (original). The method of claim 50 wherein D⁺ is a protonated form of trimethyl amine, triethyl amine, triisopropyl amine, tributyl amine, triamyl amine, isopropyldimethyl amine, t-butyldimethyl amine, diisopropylethyl amine, N,N,N',N'-tetramethyl-1,2-diaminoethane, DBU, N-methylmorpholine, N-methylpyrrolidine, N-methylpiperidine, N,N'-dimethylpiperazine, N-ethylpyrrolidine, N-ethylpiperidine, N,N'-diethylpiperazine, 1,5-diazabicyclo[4.3.0]non-5-ene, 1,4-diazabicyclo[2.2.2]octane, or 1,5,7-triazabicyclo[4.4.0]dec-5ene, 2,4,6-collidine, 2,6-lutidine, pyridine, 2-methylpyridine, 2,6-diethylpyridine, 2,6-di(t-butyl)pyridine, 4-methyl-2,6-di(t-butyl)pyridine, or 2,4,6-tri(t-butyl)pyridine, 4-dimethylaminopyridine, or N,N,N'N'-tetramethylguanidine, or tetramethylammonium, tetraethylammonium, tetrapropylammonium, tetrabutylammonium, trimethyloctylammonium, or triethylbenzylammonium cation; and

E is 1H-tetrazolide anion, 4,5-dicyanoimidazolide anion, methylsulfonate anion, trifluoromethylsulfonate anion, methylphenylsulfonate anion, trifluoromethylphenylsulfonate anion,

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tetrafluoroborate anion, hexafluorophosphate anion, or trifluoroacetate anion.

93 (original). The method of claim 50 wherein Q is O; Z is O;

Pg is β-cyanoethyl, methyl, (N-methyl-N-benzoylamino)ethyl, (N-ethyl-N-benzoylamino)ethyl, 2-[N-methyl-N-(4-methoxybenzoyl)amino]ethyl, 2-(N-isopropyl-N-benzoylamino)ethyl, 2-[N-ethyl-N-(4-methoxybenzoyl)amino]ethyl, 2-[N-isopropyl-N-(4-methoxybenzoyl)amino]ethyl, 2-[N-methyl-N-(4-dimethylaminobenzoyl)amino]ethyl, 2-[N-isopropyl-N-(4-dimethylaminobenzoyl)amino]ethyl, 2-[N-isopropyl-N-(4-dimethylaminobenzoyl)amino]ethyl, 2-(thionobenzoylamino)ethyl, 3-(thionobenzoyl-amino)propyl, 2-(N-phenylthiocarbamoylamino)ethyl, 2-[(1-naphthyl)carbamoyloxy]ethyl, diphenyl-silylethyl, δ-cyanobutenyl, cyano p-xylyl, methyl-N-trifluoroacetyl ethyl or acetoxy phenoxy ethyl; and

Y is β -cyanoethyl, allyl, methyl, (N-methyl-N-benzoylamino)ethyl, (N-ethyl-N-benzoylamino)ethyl, 2-[N-methyl-N-(4-methoxybenzoyl)amino]ethyl, 2-[N-isopropyl-N-benzoylamino)ethyl, 2-[N-ethyl-N-(4-methoxybenzoyl)amino]ethyl, 2-[N-isopropyl-N-(4-methylaminobenzoyl)amino]ethyl, 2-[N-methyl-N-(4-dimethylaminobenzoyl)amino]ethyl, 2-[N-ethyl-N-(4-dimethylaminobenzoyl)amino]ethyl, 2-[N-isopropyl-N-(4-dimethylaminobenzoyl)amino]ethyl, 2-(thionobenzoylamino)ethyl, 3-(thionobenzoylamino)propyl, 2-(N-phenylthiocarbamoylamino)ethyl, 2-[N-naphthyl)carbamoyloxy]ethyl, diphenylsilylethyl, δ -cyanobutenyl, cyano p-xylyl, methyl-N-trifluoroacetyl ethyl, acetoxy phenoxy ethyl, or a negative charge.

94 (original). The method of claim 48 wherein: said neutralizing agent is a salt of formula D⁺E⁻;

E⁻ is a tetrazolide anion:

D⁺ is a protonated form of a mono-, di- or trialkyl pyridine that is optionally substituted with an amino group;

Q is O;

Z is O;

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 R_4 and R_5 are each diisopropyl, or R_4 and R_5 together with the nitrogen atom to which they are attached form morpholine;

Pg is β -cyanoethyl, methyl, diphenylsilylethyl, δ -cyanobutenyl, cyano p-xylyl, methyl-N-trifluoroacetyl ethyl or acetoxy phenoxy ethyl; and

Y is β -cyanoethyl, allyl, methyl, diphenylsilylethyl, δ -cyanobutenyl, cyano p-xylyl, methyl-N-trifluoroacetyl ethyl or acetoxy phenoxy ethyl or a negative charge.

95 (original). The method of claim 94 wherein:

E is 1H-tetrazolide anion;

D⁺ is a protonated form of dimethylaminopyridine;

Pg is β -cyanoethyl, diphenylsilylethyl, δ -cyanobutenyl, cyano p-xylyl, methyl-N-trifluoroacetyl ethyl or acetoxy phenoxy ethyl; and

Y is β -cyanoethyl, allyl, diphenylsilylethyl, δ -cyanobutenyl, cyano p-xylyl, methyl-N-trifluoroacetyl ethyl, acetoxy phenoxy ethyl or a negative charge.

96 (original). A method comprising the steps of:

- (a) providing a solid support having a 5'-O-protected phosphorus-linked oligomer bound thereto, said phosphorus-linked oligomer having at least one phosphoryl internucleoside linkage that does not bear a phosphoryl protecting group;
- (b) deprotecting the 5'-hydroxyl of the 5'-O-protected phosphorus-linked oligomer with a deprotecting reagent;
- (c) washing the deprotected phosphorus-linked oligomer on the solid support with a solution containing a neutralizing agent;
- (d) reacting the deprotected 5'-hydroxyl with an 5'-protected nucleoside phosphoramidite to produce a phosphite triester linkage therebetween; and
 - (e) oxidizing or sulfurizing the covalent linkage to form a phosphodiester, phosphorothioate,

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phosphorodithioate or H-phosphonate linkage; and

optionally repeating steps b through e at least once for subsequent couplings of additional nucleoside phosphoramidites;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, a guanidine, or a salt of formula D⁺E wherein:

D⁺ is a quaternary tetraalkylammonium cation, or a protonated form of an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, or a guanidine; and

E⁻ is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

97 (original). A method comprising the steps of:

- (a) providing a solid support having a 5'-O-protected phosphorus-linked oligomer bound thereto, said phosphorus-linked oligomer having at least one phosphoryl internucleoside linkage that does not bear a phosphoryl protecting group;
- (b) deprotecting the 5'-hydroxyl of the 5'-O-protected phosphorus-linked oligomer with a deprotecting reagent to form a support bound 5'-deprotected phosphorus-linked oligomer;
 - (c) optionally washing the deprotected phosphorus-linked oligomer on the solid support;
- (d) contacting the support bound 5'-deprotected phosphorus-linked oligomer with a solution comprising a 5'-protected nucleoside phosphoramidite to produce a phosphite triester linkage therebetween, wherein said solution further comprises a neutralizing agent; and
- (e) oxidizing or sulfurizing the phosphite triester linkage to form a phosphodiester, phosphorothioate, phosphorodithioate or H-phosphonate linkage; and

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optionally repeating steps b through e at least once for subsequent couplings of additional nucleoside phosphoramidites;

wherein said neutralizing agent is:

an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, a guanidine, or a salt of formula D⁺E⁻ wherein:

D⁺ is a quaternary tetraalkylammonium cation, or a protonated form of an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, or a guanidine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

98 (currently amended). A composition comprising a 5'-protected nucleoside phosphoramidite and a salt of formula D⁺E⁻ wherein:

D⁺ is a quaternary tetraalkylammonium eation, or a protonated form of an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, or a guanidine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion.

99 (currently amended). The composition of claim-98 A composition comprising a 5'protected nucleoside phosphoramidite and a salt of formula D⁺E⁻ wherein:

E is a tetrazolide anion; and

D⁺ is a protonated form of a mono-, di- or trialkyl pyridine that is optionally substituted

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with an amino group.

100 (currently amended). The composition of claim 98 A composition comprising a 5'protected nucleoside phosphoramidite and a salt of formula D⁺E⁻ wherein:

E- is 1H-tetrazolide anion; and

D⁺ is a protonated form of dimethylaminopyridine.

101 (currently amended). The composition of claim 98 further comprising A composition comprising:

-- a 5'-protected nucleoside phosphoramidite; and

-- a salt of formula D+E wherein:

D⁺ is a quaternary tetraalkylammonium cation, or a protonated form of an aliphatic amine, an aliphatic heterocyclic amine, an aromatic amine, an aromatic heterocyclic amine, or a guanidine; and

E is a tetrazolide anion, 4,5-dicyanoimidazolide anion, a substituted or unsubstituted alkylsulfonate anion, a substituted or unsubstituted arylsulfonate anion, tetrafluoroborate anion, hexafluorophosphate anion, or a trihaloacetate anion; and

-- a solid support having a 5'-O-protected phosphorus-linked oligomer bound thereto, said phosphorus-linked oligomer having at least one phosphoryl internucleoside linkage that does not bear a phosphoryl protecting group.

102 (canceled)

103 (original). The composition of claim 100 further comprising a solid support having a 5'-O-protected phosphorus-linked oligomer bound thereto, said phosphorus-linked oligomer having at least one phosphoryl internucleoside linkage that does not bear a phosphoryl protecting group.

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104. (Previously added) The method of claim 50 wherein D+ is a protonated form of alkyl, alkenyl or alkynyl amine having from one to about 20 carbons, an aliphatic heterocyclic amine, an aromatic heterocyclic amine, or a guanidine.

105. (Canceled)